

### Simple calculations regarding contribution of pathways

Suppose we have four pathways, each contributing  $p_1$ ,  $p_2$ ,  $p_3$  and  $p_4$  percent to the dose (or risk)

$$p_1 = 100 * \text{dose}_1 / \text{dose}$$

$$p_2 = 100 * \text{dose}_2 / \text{dose}$$

$p_3$ ,  $p_4$ , etc

Now suppose one of the pathway doses, say  $\text{dose}_1$ , is underestimated (overestimated) by some amount  $u_1$ . Then a new overall dose, and percent contributions can be calculated.

$$\text{dose}' = \text{dose} + u_1$$

$$p_1' = 100 * (\text{dose}_1 + u_1) / (\text{dose} + u_1)$$

$$p_2' = 100 * \text{dose}_2 / (\text{dose} + u_1)$$

etc.

By examination, we see the obvious –  $p_1'$  is greater than  $p_1$ , and the other  $p$ 's are decreased proportionately.

Since the RSAL is inversely proportional to dose (or risk), the net effect of an increase in a pathway contribution is a decrease in the RSAL proportioned inversely to the overall dose (or risk) increase.

$$\text{RSAL}' = \text{RSAL} * \text{dose} / (\text{dose} + u_1)$$

EXAMPLE: Assume  $p_1$  is 52%,  $p_2$ ,  $p_3$  and  $p_4$  are each 16%, corresponding for simplicity to doses of 52 and 16 units respectively. Suppose now that  $\text{dose}_1$  has been underestimated by 26 units, or 50%.

$$p_1' = 100 * (52+26) / (100+26)$$

$$p_1' = 62\%, \text{ an increase of } 10\%$$

and

$$\text{RSAL}' = \text{RSAL} * 100 / 126$$

$$\text{RSAL}' = \text{RSAL} * 79\%$$

In this example, an underestimate of 50% in the major contributing pathway has resulted in approximately a 20% overestimate of the RSAL. An understanding of this relationship is very important to interpretation of uncertainties in the RSAL calculations.

ADMIN RECORD

SW-A-006051